## REMARKS

Entry of the foregoing, reexamination and further and favorable reconsideration of the subject application in light of the following remarks, pursuant to and consistent with 37 C.F.R. § 1.129(a), are respectfully requested.

By the present amendment, independent claims 46 and 52 have been amended. Support for the amendment of claims 46 and 52 may be found, at the very least, at page 14, lines 5-19, and especially lines 14-16. No new matter is thus being added by this amendment.

Applicant respectfully requests a personal interview after the Examiner's initial review of this amendment, but prior to issuance of a further official action..

Turning now to the Official Action, the Examiner's rejections of the claims have been obviated by the present amendment. None of the references of record contemplate administration to an animal, and thus say nothing about the special problems associated with the preparation of such liposome vesicles. Moreover, the comparative data requested by the Examiner is contained in Example 3 at pages 18-19.

In view of the above, the Examiner is respectfully requested to withdraw the rejections of the claims.

Further and favorable action in the form of a Notice of Allowance is respectfully requested. Such action is believed to be in order.

In the event that there are any questions relating to this Amendment, or the application in general, prior to the requested personal interview it would be appreciated if the Examiner would contact the undersigned attorney by telephone so that prosecution is expedited. In any event, the undersigned awaits being contacted for the requested personal interview.

Respectfully submitted,

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Date: December 17, 2001

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**Attachment to Amendment and Reply**
**Pursuant to 37 C.F.R. §1.129(a) dated December 17, 2001**

**Marked-up Claims 46 & 52**

46. (Amended) A method for preparing a stable liposome vesicle-entrapped chemical species which comprises the steps of:

(a) forming liposomes in:

(1) an aqueous medium containing an acid which is substantially impermeable through the vesicle to give an acidic liposome-containing aqueous medium in which the acid is present in the internal and external liposome phases; or

(2) an aqueous medium containing a base which is substantially impermeable through the vesicle to give an basic liposome-containing aqueous medium in which the base is present in the internal and external liposome phases;

(b) adding:

(1) to the thus-obtained acid liposome-containing aqueous medium a permanently charged, chargeable, or pH titratable chemical species which is a cationic chemical species, or

(2) to the thus-obtained acid liposome-containing aqueous medium a permanently charged,

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**Marked-up Claims 46 & 52**

chargeable, or pH titratable chemical species which is an anionic chemical species; and

(c) adding to the external liposome phase:

(1) a base to thereby induce the cationic chemical species to pass into the liposomes' internal acidic aqueous phase, or

(2) an acid to thereby induce the anionic chemical species to pass into the liposomes' internal basic aqueous phase;

wherein said cationic chemical species or said anionic chemical species is accumulated and entrapped within said liposome to produce a stable liposome vesicle-entrapped chemical species, said stability being such that after administration to an animal the chemical species is carried to its destination by the liposome vesicle before significant leakage occurs.

52. (Amended) A method of preparing a stable liposome vesicle-entrapped chemical species, which method comprises:

(a) forming liposomes in:

(1) an aqueous medium containing an acid which is substantially impermeable through the vesicle to give an acidic liposome-containing aqueous medium

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**Marked-up Claims 46 & 52**

in which the acid is present in the internal and external liposome phases; or

(2) an aqueous medium containing a base which is substantially impermeable through the vesicle to give an basic liposome-containing aqueous medium in which the base is present in the internal and external liposome phases;

(b) adding:

(1) to the thus-obtained acid liposome-containing aqueous medium a permanently charged, chargeable, or pH titratable chemical species which is a cationic chemical species, or

(2) to the thus-obtained acid liposome-containing aqueous medium a permanently charged, chargeable, or pH titratable chemical species which is an anionic chemical species; and

(c) adding to the external liposome phase:

(1) a base in an amount effective to create a pH gradient between the external liposome phase and the internal liposome phase to thereby induce the

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**Marked-up Claims 46 & 52**

cationic chemical species to pass into the liposomes' internal acidic aqueous phase, or

(2) an acid in an amount effective to create a pH gradient between the external liposome phase and the internal liposome phase to thereby induce the anionic chemical species to pass into the liposomes' internal basic aqueous phase;

wherein said cationic chemical species or said anionic chemical species is accumulated and entrapped within said liposome to produce a stable liposome vesicle-entrapped chemical species, said stability being such that after administration to an animal the chemical species is carried to its destination by the liposome vesicle before significant leakage occurs.